

## (12) INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(19) World Intellectual Property  
Organization  
International Bureau



(43) International Publication Date  
28 October 2004 (28.10.2004)

PCT

(10) International Publication Number  
**WO 2004/091480 A2**

(51) International Patent Classification<sup>7</sup>: **A61K**  
(21) International Application Number:  
PCT/US2004/010626  
(22) International Filing Date: 8 April 2004 (08.04.2004)  
(25) Filing Language: English  
(26) Publication Language: English  
(30) Priority Data:  
60/461,471 9 April 2003 (09.04.2003) US

(71) Applicant (for all designated States except US): **EX-ELIXIS, INC.** [US/US]; 170 Harbor Way||P.O. Box 511, South San Francisco, CA 94083-0511 (US).

(72) Inventors; and

(75) Inventors/Applicants (for US only): **CHEN, Jeff** [US/US]; 140 So. Van Ness Avenue, Apt. 407, San Francisco, CA 94103 (US). **DALRYMPLE, Lisa** [US/US]; 387 7th Avenue, San Francisco, CA 94121 (US). **EPSHTEYN, Sergery** [US/US]; 2922 Southwycke Terrace, Fremont, CA 94536 (US). **FORSYTH, Timothy** [US/US]; 1928 Wingate Way, Hayward, CA 94541 (US). **HUYNH, Tai** [US/US]; 1530 6th Avenue #2, Oakland, CA 94606 (US). **LEAHY, James** [US/US]; 1185 Camellia Court, San Leandro, CA 94110 (US). **MANN, Grace** [US/US]; 231 Calippe Court, Brisbane, CA 94005 (US). **MANN, Larry, W.** [US/US]; 780 Bair Island Road, Apt. 304, Redwood City, CA 94063 (US). **RIDGWAY, Brian** [US/US]; 2403 Palmer Avenue, Belmont, CA 94002 (US). **SANGALANG, Joan, C.** [US/US]; 679 Fairmont Avenue,

#5, Mountain View, CA 94041 (US). **TAKEUCHI, Craig** [CA/US]; 1090 Carolan Avenue #302, Burlingame, CA 94010 (US).

(74) Agents: **GRIEDEL, Brian et al.**; Exelixis, Inc., 170 Harbor Way, P.O. Box 511, South San Francisco, CA 94083-0511 (US).

(81) Designated States (unless otherwise indicated, for every kind of national protection available): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW.

(84) Designated States (unless otherwise indicated, for every kind of regional protection available): ARIPO (BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European (AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

## Published:

— without international search report and to be republished upon receipt of that report

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

(54) Title: **TIE-2 MODULATORS AND METHODS OF USE**

(57) Abstract: The present invention provides compounds for modulating protein kinase enzymatic activity for modulating cellular activities such as proliferation, differentiation, programmed cell death, migration and chemoinvasion. Compounds of the invention inhibit, regulate and/or modulate kinases, particularly Tie-2. Methods of using the compounds and pharmaceutical compositions thereof to treat kinase-dependent diseases and conditions are also an aspect of the invention.